

FIG. 1A





VALIDATION - PHASE 1

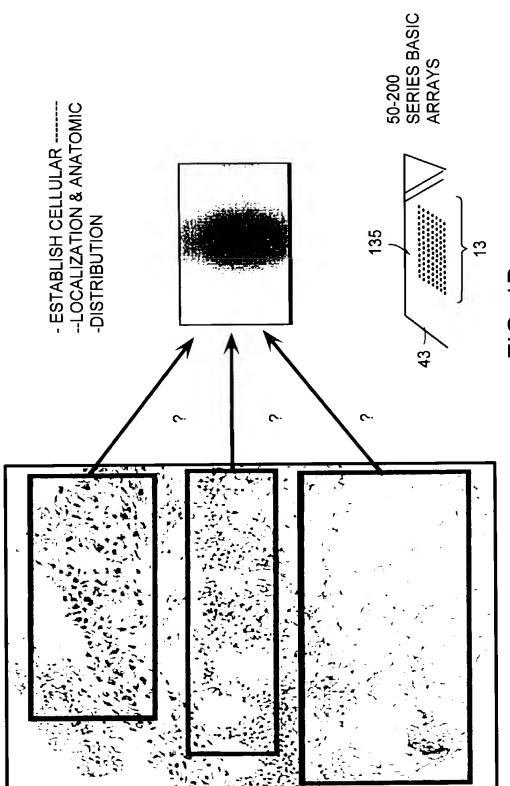
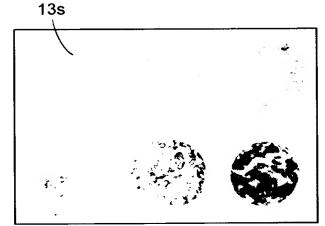


FIG. 1B



FIG. 2A



SIX TISSUE ELEMENTS FROM A PathWorkTM BreastArray BR50 STAINED WITH CK7 SHOWING SELECTIVE STAINING (400X). TISSUE **ELEMENTS ARE 0.6mm DIAMETER AND SPACED** 0.3mm APART.

FIG. 2C

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FIG. 2B



43a__ 43b-

FIG. 2D

SAMPLE DATA SHEET INFORMATION (FOR ONE TISSUE ELEMENT, LOCATED AT A1) PROVIDED WITH EVERY PathWorks™ Tissue MicroArray

BR50 CAT.#

07 LOT#

COORDINATES: A1a MITOSES: 1 **BREAST** PT: 1

ORGAN: BREAST CANCER, DUCTAL 0 pN: **CATEGORY**:

F 0 SEX: LN POS

67 23 LN ALL AGE:

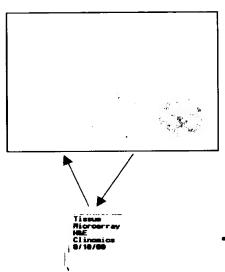
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FIG. 2E





TISSUE MicroArrays

- DEVELOPMENT OF POPULATION DATABASES
- HIGH-THROUGHPUT IN SITU ANALYSIS OF DNA, RNA AND PROTEIN
- DISCOVERY OF DIAGNOSTIC AND PROGNOSTIC CORRELATIONS
- TARGET PRIORITIZATION

FIG. 3

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CLIN		ords Tools Window	(a) (c)	Specimen		Specimen Data	Received Date 10/15/1999 Metastasis Site Lung RadiationTherapy Yes SurgicalResection Yes	177.
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FIG. 4A	Clinom	Elle	<u>}</u>		Path	Patient	Medication CMF Chemother Chemother PrimarySit (Breast (ClinomicsID 000002841 000007489 000007491 000007493 4 Corm View

DATA ENTRY

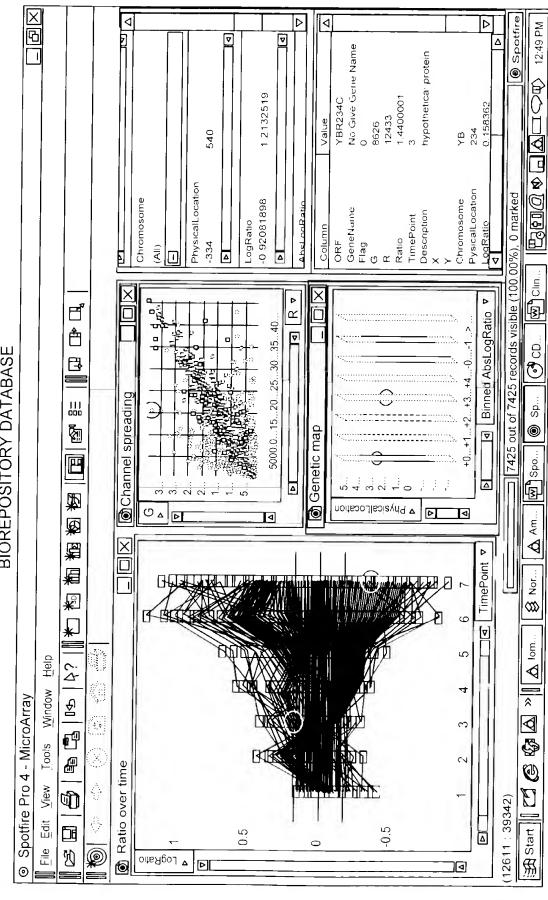


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SORTING AND SELECTION OF PATIENT POPULATIONS

CLUSTER ANALYSIS

DEVELOPMENT OF EXPRESSION PROFILES VS MEDICAL DATA

FIG. 4C



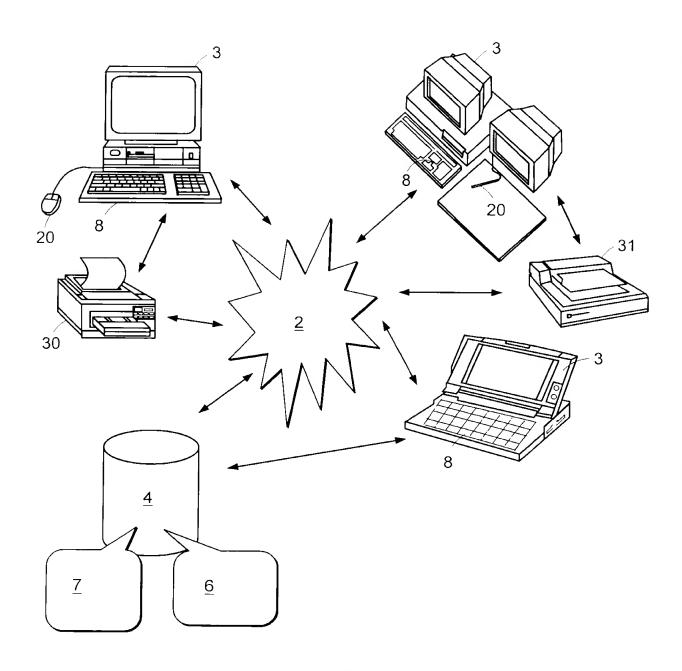
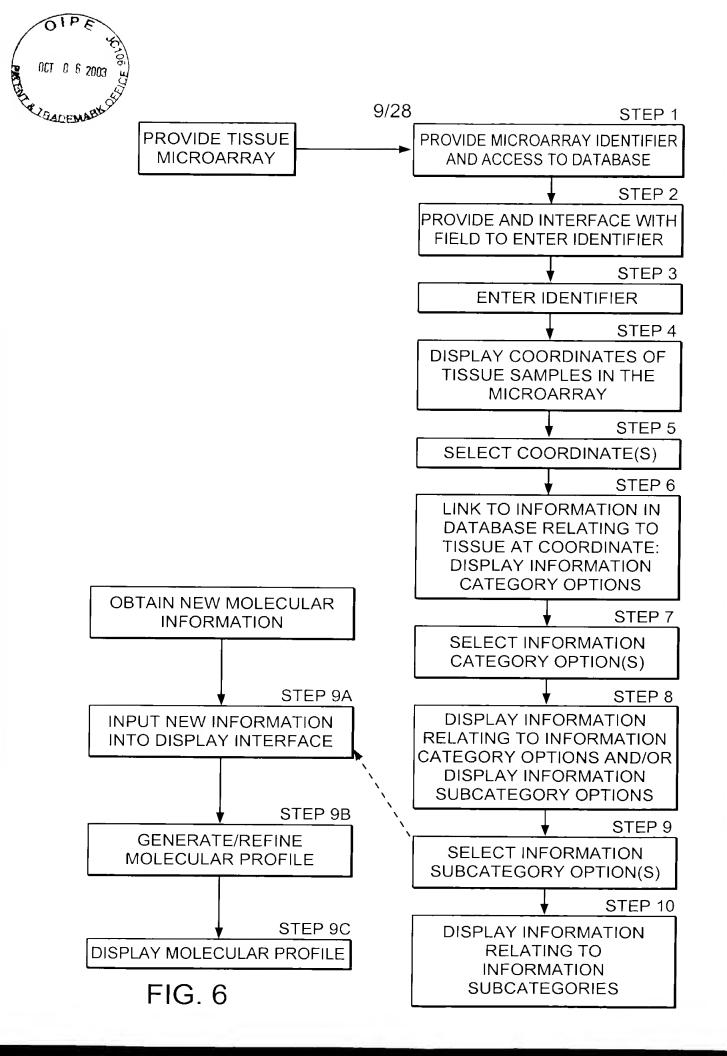
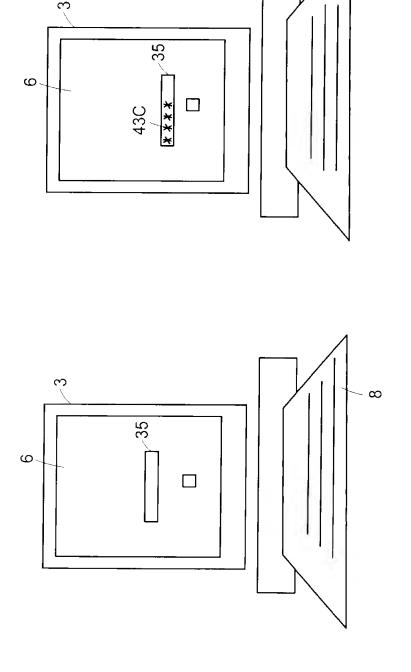


FIG. 5







:IG: 7A

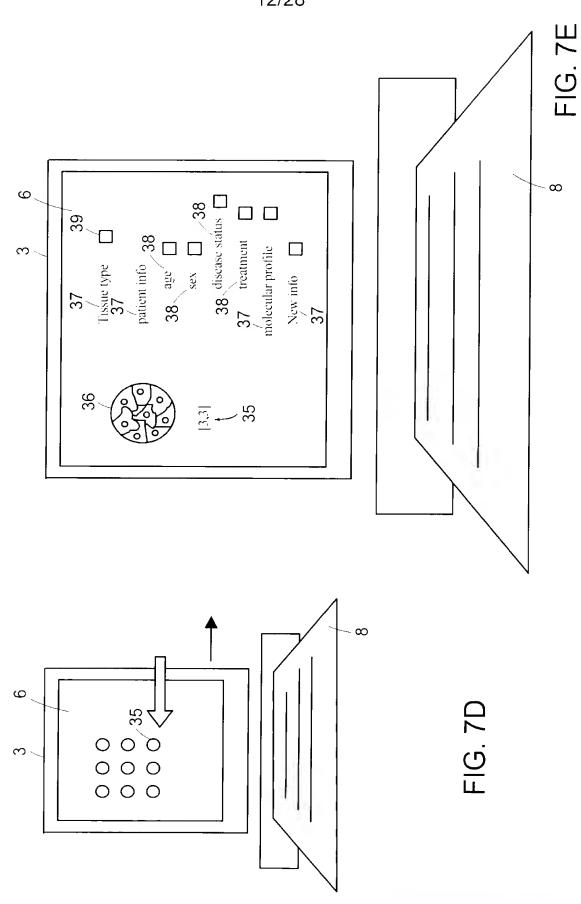
FIG 7B

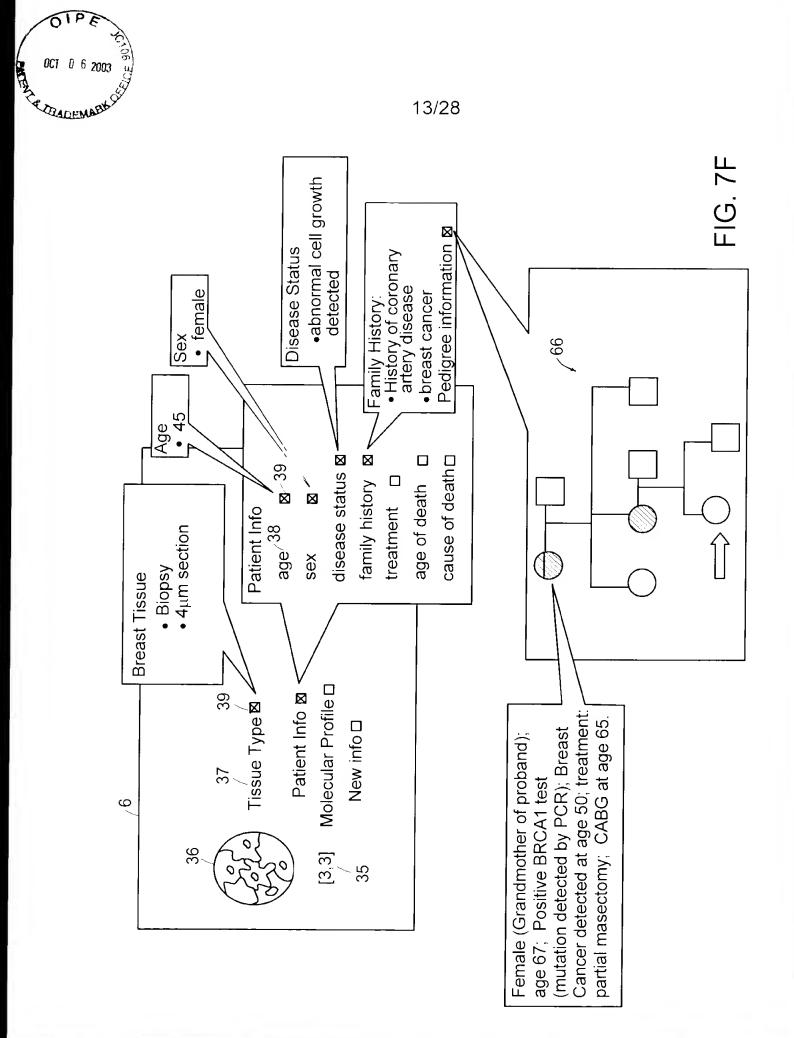


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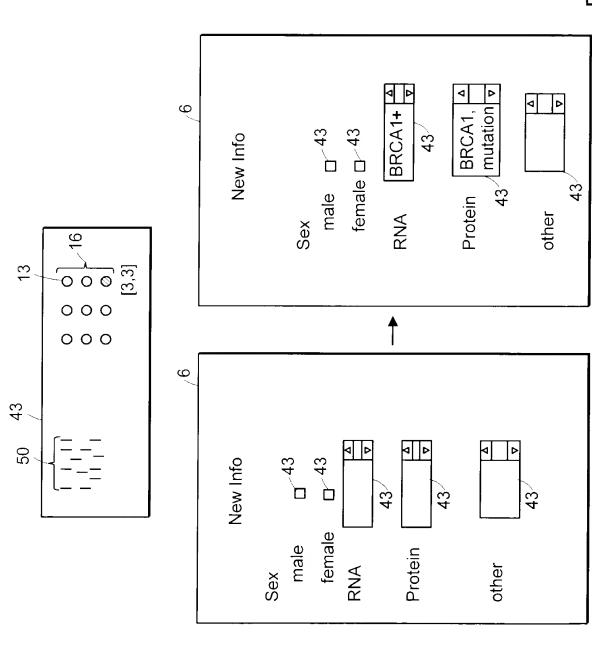
FIG. 7C

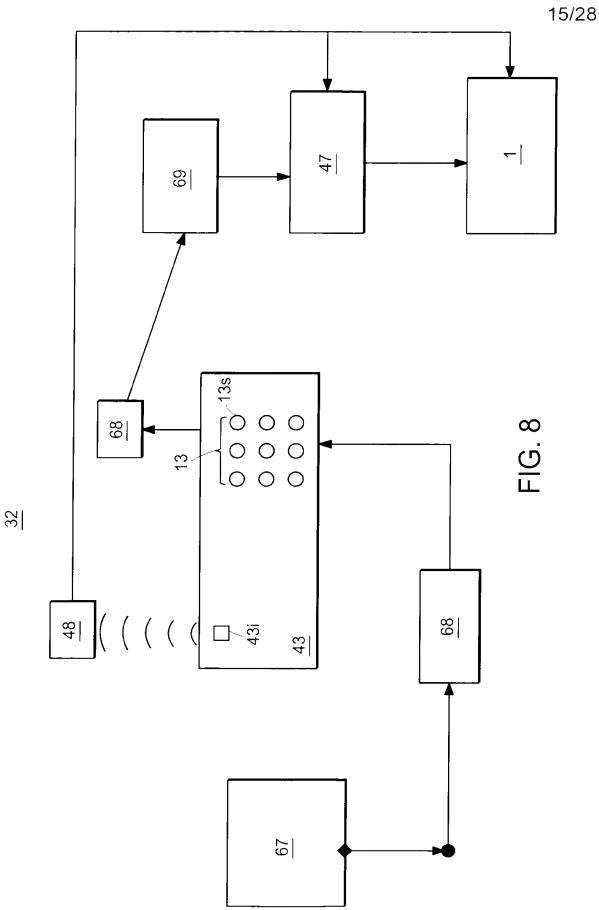
















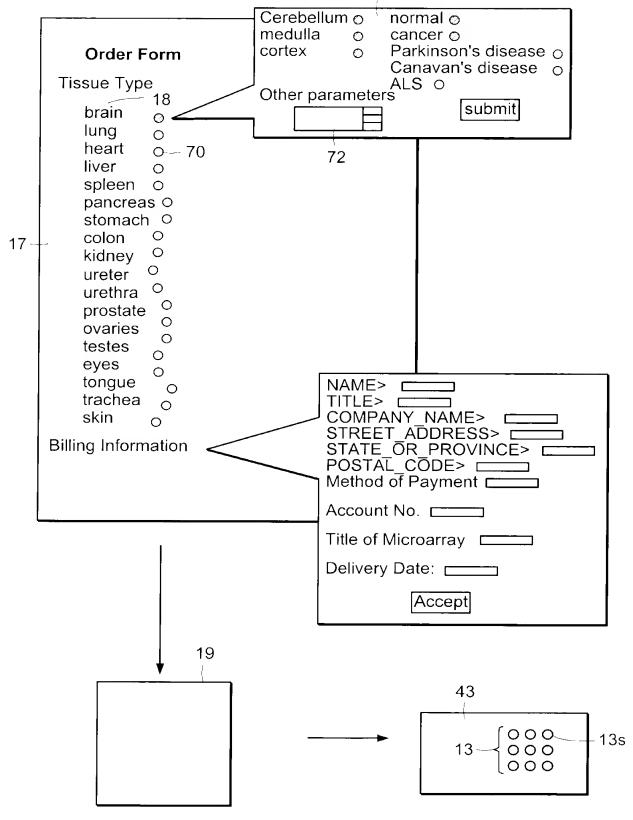
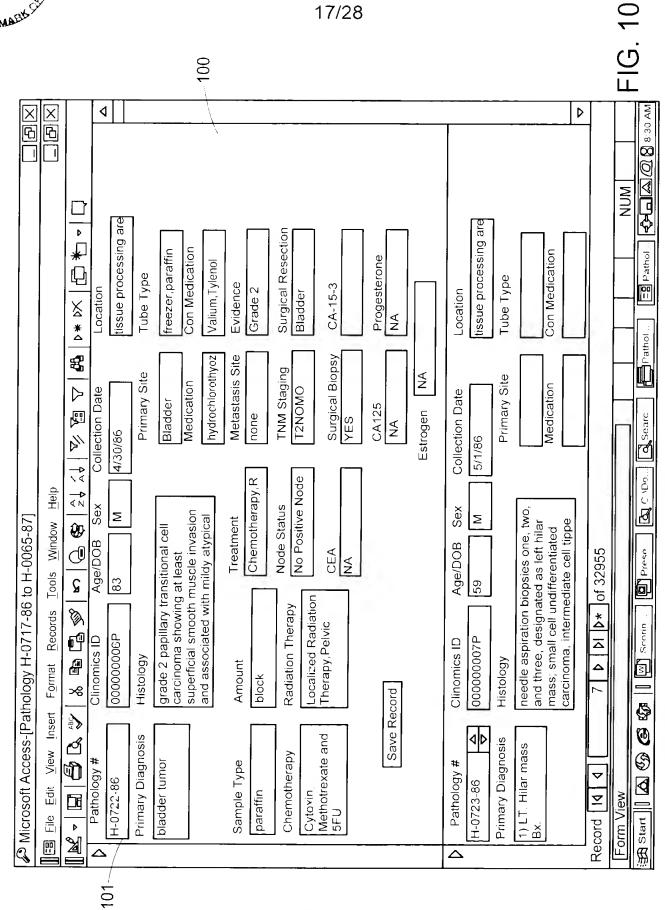


FIG. 9







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FIG. 11B



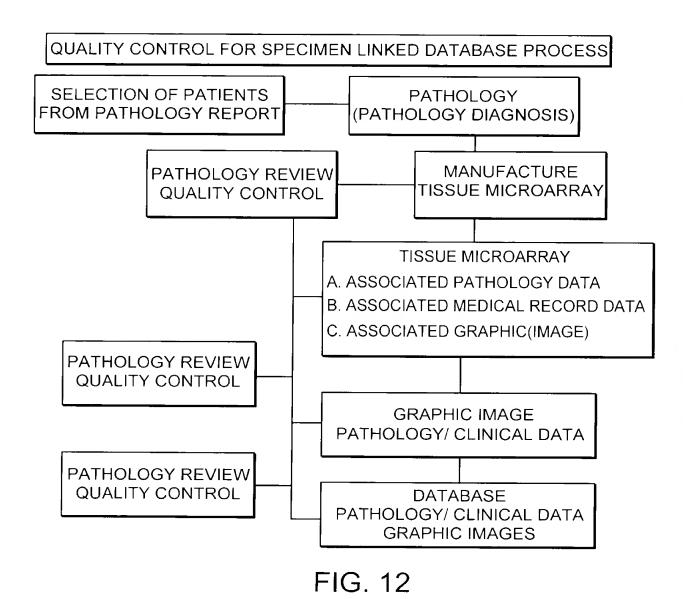


FIG. 13A FIG. 13B

FIG. 13

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T.02	SKIN	T.41	ARTERY, GENERAL	1.77	PROSTATE	T.Y1	T.Y1 TRUNK PERINEUM
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							RETROPERITONEUM
90.T	BONE MARROW	T.44	PULM,ARTERY	T.80	VULVA, LABIUM, CLITORIS T. Y6 PELVIS	T.Y6	PELVIS
-					BARTOLINS GLAND	T.YX	T.YX PERITONEAL FLUID
T.07	SPLEEN	T.45	VERT, CAROTID, CEREBR, TEMPOR.	T.81	VAGINA, HYMEN	T.Y1	T.Y1 SYNOVIAL FLUID
			BASILOR ARTERIES			T.2X	T.2X UPPER RESP FLDS
T.08	LYMPH NODES	T.46	ILIAC, RENAL, MESSENTERIC ARTERY	T.82	UTERUS	T.2Y	LOWER RESP FLDS
T.09	LYMPHATICS	T.47	POPLITEAL ARTERY	T.83	CERVIX	T.6X	T.6X GI FLDS, SPACES
1.11	BONE	T.48	VEIN	T.84	ENDOMETRIUM	T.6Y	GI CONTENTS
T.12	SYNOVIUM,JOINT	T.49	VEIN	T.85	MYOMETRIUM	T.7X	T.7X URINE TRA.FLO. SPACE
T.13	MUSCLE	T.50	DIGESTIVE SYSTEM	T.86	FALLOPIAN TUBE	T.7Y	T.7Y MALE GENT FLD, SPACE
T.15	CARTILADGE	T.51	MOUTH, PALATE, ORAL	T.87	OVARY	¥8.⊤	FEM. GENT FLD. SPACE
T.16	BURCA	T.52	LIP	T.88	PLACENTA, UMBCORD T.8Y PLACENTA, FETAL FLD	T.8Y	PLACENTA, FETAL FLD
T.17	TENDON	T.53	TONGUE	T.89	FETUS	T.XO	T. XO NERVOUS SYSTEM
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FIG. 13A

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FIG. 13E

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CARCINOMA, PAPILLARY CARCINOMA, SMALL CELL M.8043 CARCINOSARCOMA M.8983 CARCINOID, MALIGNANT TUMOR CARCINOMA, UNDIFFERENTIATED METS CARCINOMA, UNDIFFERENTIATED M.8023 CHORIOCARCINOMA M.9103 CARCINOID TUMOR M.8241 CHOLANGIOCARCINOMA M.8163 CHONDROSARCOMA M.9223 CHONDROSARCOMA M.9230 CRANIOPHARYNGIOMA M.8231 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8203 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CARCINOMA, SMALL CELL CARCINOSARCOMA CARCINOID, MALIGNANT TUMOR CARCINOMA, UNDIFFERENTIATED METS M.8023 CARCINOMA, UNDIFFERENTIATED M.8023 CHORIOCARCINOMA CARCINOID TUMOR CARCINOID TUMOR CHOLANGIOCARCINOMA CHONDROSARCOMA CHONDROSARCOMA CHONDROBLASTOMA CRANIOPHARYNGIOMA M.8230 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CARCINOSARCOMA CARCINOID, MALIGNANT TUMOR CARCINOMA, UNDIFFERENTIATED METS M.8023 CARCINOMA, UNDIFFERENTIATED M.8023 CHORIOCARCINOMA M.9103 CARCINOID TUMOR M.8241 CHOLANGIOCARCINOMA M.8163 CHONDROSARCOMA M.9223 CHONDROBLASTOMA M.9230 CRANIOPHARYNGIOMA M.8203 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CARCINOID, MALIGNANT TUMOR M.8243 CARCINOMA, UNDIFFERENTIATED METS M.8023 CHORIOCARCINOMA M.9103 CARCINOID TUMOR M.8241 CHOLANGIOCARCINOMA M.8163 CHONDROSARCOMA M.9223 CHONDROBLASTOMA M.9230 CRANIOPHARYNGIOMA M.8203 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8203 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CARCINOMA, UNDIFFERENTIATED METS M.8023 CARCINOMA, UNDIFFERENTIATED M.8023 CHORIOCARCINOMA M.9103 CARCINOID TUMOR M.8241 CHOLANGIOCARCINOMA M.8163 CHONDROSARCOMA M.9223 CHONDROBLASTOMA M.9230 CRANIOPHARYNGIOMA M.9230 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8203 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503	CARCINOID, MALIGNANT TUMOR	
CARCINOMA, UNDIFFERENTIATED M.8023 CHORIOCARCINOMA M.9103 CARCINOID TUMOR M.8241 CHOLANGIOCARCINOMA M.8163 CHONDROSARCOMA M.9223 CHONDROBLASTOMA M.9230 CRANIOPHARYNGIOMA M.9351 CYLINDROMA M.8203 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503	CARCINOMA, UNDIFFERENTIATED METS	M 8023
CHORIOCARCINOMA CARCINOID TUMOR CHOLANGIOCARCINOMA CHONDROSARCOMA CHONDROBLASTOMA CRANIOPHARYNGIOMA CYLINDROMA CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY DERMATOFIBROSARCOMA DUCTAL, INFILTRATING CARCINOMA M.9103 M.8163 M.9230 M.8203 M.8463 M.8463		
CARCINOID TUMOR CHOLANGIOCARCINOMA M.8163 CHONDROSARCOMA M.9223 CHONDROBLASTOMA M.9230 CRANIOPHARYNGIOMA M.9351 CYLINDROMA M.8203 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CHOLANGIOCARCINOMA M.8163 CHONDROSARCOMA M.9223 CHONDROBLASTOMA M.9230 CRANIOPHARYNGIOMA M.9351 CYLINDROMA M.8203 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CHONDROSARCOMA M.9233 CHONDROBLASTOMA M.9230 CRANIOPHARYNGIOMA M.9351 CYLINDROMA M.8203 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CHONDROBLASTOMA CRANIOPHARYNGIOMA M.9230 CYLINDROMA M.8203 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA DYSPLASIA DUCTAL, INFILTRATING CARCINOMA M.8503	CHONDROSARCOMA	
CRANIOPHARYNGIOMA M.9351 CYLINDROMA M.8203 CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CYSTADENOCARCINOMA PAPILLARY SEROUS M.8463 CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
CYSTADENOCARCINOMA, PAPILLARY M.8453 DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
DERMATOFIBROSARCOMA M.8823 DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
DYSPLASIA M.7600 DUCTAL, INFILTRATING CARCINOMA M.8503		
DUCTAL, INFILTRATING CARCINOMA M.8503		
	EPENDYMOMA	M.9391

FIG. 14A

001 0 0 30	(A)		
OCT 0 6 20	103 94 	20	
132	24/2 EPIDERMOID CARCINOMA EMBRYONAL CELL CARCINOMA EPITHELOMA	28	
MADEN	EPIDERMOID CARCINOMA	M.8073	
	EMBRYONAL CELL CARCINOMA	M.9073	
	EPITHELOMA	M.8110	
	EWINGS SARCOMA	M.9263	
	FIBROSARCOMA	M.8823	
	FOLLICULAR CARCINOMA	M.8333	
	GIANT CELL SARCOMA	M.8803	
	GIANT CELL TUMOR	M.9040	
1	GRANULOSA CELL TUMOR	M.8621	
	GLIOBLASTOMA, MULTIFORME	M.9443	
	GLIOMA	M 9383	
	GIANT CELL CARCINOMA	M.8033	
	HEMANGIOBLASTOMA	M.9161	
	HEPATOBLASTOMA	M 8973	
	HISTOCYTOMA	M.8830	
	HODGKINS	M.9653	
	HODGKINS SARCOMA	M.9683	
	HURTLE CELL CARCINOMA	M.8363	
	ISLET CELL CARCINOMA	M 8153	
	KAPOSI SARCOMA	M.9143	
	LEUKEMIA, ACUTE MYELOGENOUS	M.9865	
	LEUKEMIA, LYMPHOMA	M.9826	
	LEUKEMIA	M 9807	
	LEUKEMIA, CHRONIC LYMPOMA	M 9827	
	LEIOMYOSARCOMA	M.8893	
	LYMPHOMA, LYMPHOCYTIC	M.9633	
	LYMPHOMA, LYMPHOCYTIC	M.9593	
	LIMPHOMA, GIANT FOLLICULAR	M.9693	
a P	LYMPHOSARCOMA, LYMPHOCYTIC	M 9623	
	LYMPHOMA, MALIGNANT	M.9593	
	LYMPHOMA, HISTIOCYSTIC	M.9643	
	LYMPHERITHELIOMA	M.8083	
	LYMPHOMA, T-CELL	M.9703	
	LYMPHOSARCOMA, LYMPHOCYTIC	M.9613	
	LIPOSARCOMA	M 8853	.
	LYMPHOMA, STEM CELL	M 9603	
	MALIGNANT MELANOMA SPINDLE CEI		
	MENINGIOMA METASTATIC CARCINOMA	M 9530	
	METASTATIC CARCINOMA MALIGNANT TUMOR	M.8016 M.8003	
	METASTATIC MALIGNANT TUMOR		
	MEDULLOBLASTOMA	M 8006	_
	MALIGNANT MELANOMA	M 9473 M 8723	
	MALIGNANT CELLS	M.6972	
	MYOBLASTOMA	M 9370	
	MUCOEPIDERMOID CARCINOMA	M.8433	
	MALIGNANT CARCINOID TUMOR	M 8243	
	MALIGNANT CARCINOID TOMOR MALIGNANT MESENSHYMOMA	M.8993	
	MESOTHELIOMA	M 9051	
	MYELOMA	M 9733	-
	METASTATIC MELANOMA	M 8733	
		= [M 0733] = [C 14P	L
		11 7 4 4 1 3	

FIG. 14B



MALIGNANT GRANULOSA CELL TUMOR	M.8623
MALIGNANT PHEOCHROMOCYTOMA	M.8703
MALIGNANT MESOTHELIOMA	M.9053
METASTATIC LEIOMYOSARCOMA	M.8893
MESODERMAL TUMOR	M.8953
MIXED TUMOR	M.8940
NEUROBLASTOMA, NEUROEPITHELIOMA	1
NECROTIC TUMOR	M.5443
NEUROBLASTOMA, NEUROEPITHELIOMA	
OLIGODENDROGLIOMA	M.9453
OAT CELL CARCINOMA	M.8043
OSTEOBLASTOMA	M.9200
OSTEOMA	M.9180
OSTEOSARCOMA	M.9183
OSTEOIDOSTEOMA	M.9190
OSTEOCHONDROMA	M.9210
PAPILLARY CARCINOMA	M.8053
PLASMA CELL MYELOMA	M.9733
PAPILLARY METASTATIC CYATADENOCARCINOMA	M.8463
PAPILLARY TRANSITIONAL CELL CARCINOMA	M.8133
RENAL CELL CARCINOMA	M.2485
RETICULUM CELL SARCOMA	M.9643
RENAL CLEAR CELL CARCINOMA	M.8313
SQUAMOUS CELL CARCINOMA	M.8073
SARCOMA, RETICULAR CELL	M.8316
SQUAMOUS EPITHELIAL TUMOR	M.8070
SQUAMOUS CELL CARCINOMA IN-SITU	M.8072
SARCOMA, UNDIFFERENTIATED GIANT CELL	M.8803
SARCOMA, RETICULAR CELL	M.9643
SPINDLE CELL CARCINOMA	M.8033
SEMINOMA	M.9063
STROMAL SARCOMA	M.8933
TUMOR	M.8001
TUMORLET	M.8043
THYMOMA MALIGNANT	M.8583
TRANSITIONAL CELL CARCINOMA	M.8123
TERATOMA	M.9081
VERRICOUS SQUAMOUS CARCINOMA	M.8053

FIG. 14C

FIG. 15A

FIG. 15B

FIG. 15C

FIG. 15

2	6	10)	R
_	U	/ 4	_	U

OCT O 5 2003 LA		26/28	
MADEMARK	METHO FFPE	20/20	H
	MEDICATIONS N/A	BETAMETHASONE MAGNESIUM SULFATE PITOCIN APRESOLINE MANNITOL	N/A
	291.89 291.89	293.83	293.83
	851.80 852.00 860.0	401.9 245.9 493.9 642.0 437.2 437.2 348.5	174.9 425.4 428.0
BRAIN INVENTORY CLASSIFICATIONS	MEDICAL HISTORY MULTIPLE TRAUMATIC INJURIES SUSTAINED IN A MOTOR VEHICLE ACCIDENT OPEN SKULL FRACTURES. BRAIN CONTUSIONS. SUBARACHRICID HEMORRHAGE PERICARDIAL TEAR. BILATERAL PNEUMOTHORACES. CONTUSIONS OF HEART BLOOD ALCOHOL LEVEL 95 MG/DT	PREGNANCY INDUCED HYPERTENSION THYROIDITIS ASTHMA SEVERE PRE-EDAMPSIA ANOXIC ENCEPHALOPATHY INTRACEREBRAL HEMORRHAGE HEMORRHAGE OF RIGHT PUTAMEN HYPERTENSIVE CEREBRAL EDEMA DIFFUSE ANOXIC CHANGES	BREAST CANCER IDIOOPATHIC CARDIOMYOPATHY CHRONIC CONGESTIVE HEART FAILURE. ANAPHYLACTIC SHOCK SECONDARY TO TAXOL ADMINISTRATION. SEVERE HYPOXIA AND BRAIN DEATH.
ENTO	X N N N N N N N N N N N N N N N N N N N	LL L	ш
BRAIN INVI	DOB AGE 04/08/1981 18	04/26/1972 27	12/27/1947 52
	BRAIN (ORGANIC)	BRAIN (ORGANIC)	(ORGANIC)
	AU-99-47	AU-99-61	AU-99-80

FIG. 15A

SIPE	
OCT 0 6 2003	
MADEMARK	r

FFPE						FFPE															FFPE				
N/A						N/A															N/A				
307.89					_	291.89	305.00										ì				293.83				
307.89	305.90	414.00										571.2									185.0	162.9			3313
CHRONIC PAIN	DRUG ABUSE	SEVERE CORONARY ARTERY SCLEROSIS.	OLD MYOCARDIAL INFARCTION.	CARDIOMEGALY.		SEVERE CHRONIC ALCOHOLISM.	PREVIOUS HOSPITAL ADMISSION	FOR ALCOHOL INTOXICATION AND	MAJOR DEPRESSION.	SEVERE ASCITES(4,000 ML)	HEPATOMEGALY(LIVER 1,690 GM)	SEVERE CIRRHOSIS.	MULTIPLE TRAUMATIC INJURIES	SUSTAINED IN A MOTOR VEHICLE	ACCIDENT:AVULSION OF SOFT TISSUE,	BONE AND BRAIN.	MULTIPLE COMMINUTED OPEN	FRACTURES OF THE SKUIL	ALCOHOL INTOXICATION	BLOOD ETHANOL LEVEL 252 GM %	PROSTATE CANCER	SQUAMOUS CELL CANCER IT LUNG	VENTRICULAR PERITONEAL SHUNT	SECONDARY TO PROGRESSIVE	HYDROCEPHALUS FROM BRAIN MASS
Σ		_				<u>+</u>					_		M /								Σ				
09/05/1957 42						05/10/1960 39							04/19/1962 37								01/09/2028 71				
BRAIN		(PSYCH)				BRAIN	(PSYCH)						BRAIN	(PSYCH)							BRAIN	(ORGANIC)			
AU-99-82						AU-99-84							AU-99-90								AU-99-91				

FIG. 15B

OCT OF 2003 B													28	3/2	28									
											FFPE													
						i					CCUPRIL, COLCHICINE, NAPRO													
											303.00	291.89							305.00					
	401.9					197.7	198.3	492.8			278.00	401.9	274.9			410.9		571.2		401.9	571.2		492.8	
	HYPERTENSION	HEADACHES	UNSTEADY GATE	MENTAL CONFUSION	SQUAMO	TO LIVER, SPLEEN, KIDNEYS,	AND BRAIN.	EMPHISEMA	ATHEROSCLEROSIS OF CORONARY	ARTERIES.	MORBID OBESITY	HYPERTENSION	GOUT	CRONIC PAIN SYNDROME	DEPRESSION	ACUTE MYOCARDIAL INFARCTION	ACUTE ALCOHOL INTOXICATION	MICROMODULAR CIRRHOSIS	ETHANOL ABUSE	HYPERTENSION	MICRONODULAR CIRRHOSIS	SEVERE ASCITES(5LT)	SPLENOMEGALY	EMPHISEMA
											Σ								Щ					
	_										41								20	<u> </u>				
											05/01/1958								BRAIN ###### 50					
											BRAIN	(PSYCH)							BRAIN	(PSYCH)				
											AU-99-93								AU-99-94					

FIG. 15C